

## **Supplemental Material**

### **Environmental Cadmium and Lead Exposures and Hearing Loss in US Adults: the National Health and Nutrition Examination Survey, 1999 to 2004**

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## **Table of Contents**

Supplemental Material, Table S1. General characteristics of study participants and non-participants

Supplemental Material, Table S2. Participant characteristics by hearing loss status

Supplemental Material, Table S3. Percent change (95% CIs) of hearing thresholds (dB) by contribution of different variables in a multiple linear regression

Supplemental Material, Figure S1. Graphical distribution of blood cadmium and lead with log-transformed hearing thresholds

Supplemental Material, Figure S2. Percent change (%) of hearing thresholds (dB) by blood cadmium and lead quintiles at each frequency from 0.5 kHz to 8 kHz

**Supplemental Material, Table S1.** General characteristics of study participants and non-participants (N=5263<sup>a</sup>)

Characteristic	Participants <sup>b</sup> (N=3698)	Non-Participants (N=1565)	P-value
Occupation noise exposure <sup>c</sup> ( <i>O*NET score</i> )	3.06 (± 0.02)	3.12 (± 0.02) <sup>g</sup>	0.052
Age (y)	42.06 (± 0.28)	42.09 (± 0.43)	0.956
Body mass index ( <i>wtkg/htm</i> )	28.04 (± 0.15)	28.47 (± 0.31) <sup>h</sup>	0.212
PTA Hearing Thresholds <sup>d</sup> (dB)	12.78 (± 0.24)	16.95 (± 0.40)	<.001
Hearing Loss <sup>e</sup> ( <i>PTA&gt;25dB, %</i> )	11.9	28.7	<.001
Noise notch <sup>f</sup> (%)	17.7	17.0	0.592
Sex ( <i>Male %</i> )	48.6	50.3	0.275
Race ethnicity (%)			<.001
<i>Non-Hispanic White</i>	72.5	65.7	
<i>Non-Hispanic Black</i>	10.5	11.1	
<i>Mexican American</i>	6.6	9.3	
<i>Other</i>	10.4	13.9	
Education (%)			<.001
< <i>High School</i>	16.6	21.8 <sup>i</sup>	
<i>High School</i>	25.1	24.5	
> <i>High School</i>	58.3	53.7	
Ototoxic medication ( <i>Current use %</i> )	15.9	14.8 <sup>j</sup>	0.468
Cumulative cigarette pack-years (%)			0.273
<i>Never</i>	53.7	54.3 <sup>k</sup>	
<20	33.7	31.3	
≥20	12.5	14.4	
Hypertension (%)	23.2	24.8 <sup>l</sup>	0.289
Diabetes mellitus (%)	4.1	6.4 <sup>m</sup>	8.632
Firearm noise exposure ( <i>Exposed %</i> )	7.5	9.2 <sup>n</sup>	0.188
Recreation noise exposure ( <i>Exposed %</i> )	26.0	28.5 <sup>o</sup>	0.126

Continuous variables: survey t-test, age-adjusted

Categorical variables: 2\*2 table or 2\*C table : survey X\_square (Rao-Scott Chi-Square Test)

<sup>a</sup>Subjects (N=5263) are eligible for the audiometric data

<sup>b</sup>Participants (N=3698) are the individuals having all interest variables in this study: hearing thresholds, hearing loss, age, BMI, sex, race ethnicity, ototoxic medication, cumulative cigarette packyears, hypertension, diabetes mellitus, occupational noise score, and firearm and recreation noise exposure

<sup>c</sup>Occupation noise (1 < O\*NET noise scale < 5)

<sup>d</sup>PTA at speech frequencies (Pure tone means at 0.5, 1, 2, 4 KHz)

<sup>e</sup>Hearing loss was defined as PTA at speech frequencies > 25 dB

<sup>f</sup>Noise Notch (Hearing threshold at 3,4, and/or 6 kHz is at least 10 dB greater than at 1 or 2 kHz and at least 10 dB greater than at 6 or 8 kHz.)

Non-Participants (N=1137<sup>g</sup>, 1492<sup>h</sup>, 1563<sup>i</sup>, 1559<sup>j</sup>, 1119<sup>k</sup>, 1343<sup>l</sup>, 1563<sup>m</sup>, 1560<sup>n</sup>, 1557<sup>o</sup>, and 1565 for the others) are the individuals having the variable to be compared

**Supplemental Material, Table S2.** Participant characteristics by hearing loss status (N=3698<sup>a</sup>).

Characteristic	All participants (N=3698)	Not Hearing Loss (N=3257)	Hearing Loss <sup>a</sup> (N=441)	<i>P-value</i> <sup>b</sup>
Blood Lead ( <i>ug/dL</i> )	1.94 (±0.04)	1.91 (±0.04)	2.17 (±0.09)	0.006
Blood Cadmium ( <i>ug/L</i> )	0.56 (±0.01)	0.55 (±0.01)	0.67 (±0.04)	0.003
Age ( <i>y</i> )	42.06 (±0.28)	40.45 (±0.29)	54.81 (±0.66)	<.001
Body mass index ( <i>wt/kg/ht/m</i> )	28.04 (±0.15)	28.03 (±0.15)	28.06 (±0.36)	0.941
Pure tone average hearing thresholds (dB) <sup>c</sup>	12.78 (±0.24)	10.98 (±0.16)	27.03 (±0.68)	<.001
Occupation noise exposure ( <i>O*NET score</i> ) <sup>d</sup>	3.06 (±0.02)	3.04 (±0.02)	3.25 (±0.04)	<.001
Sex ( <i>Male %</i> )	48.6	46.4	66.3	<.001
Race ethnicity (%)				<.001
<i>Non-Hispanic White</i>	72.5	71.5	80.6	
<i>Non-Hispanic Black</i>	10.5	11.2	4.5	
<i>Mexican American</i>	6.6	7.1	3.4	
<i>Other</i>	10.4	10.2	11.5	
Education (%)				0.002
< <i>High School</i>	16.6	15.1	28.3	
<i>High School</i>	25.1	24.7	28.2	
> <i>High School</i>	58.3	60.2	43.4	
Ototoxic medication ( <i>Current use %</i> )	15.9	14.8	24.3	0.002
Cumulative cigarette pack-years (%)				<.001
<i>Never</i>	53.7	55.4	40.8	
<20	33.7	34.6	27.3	
≥20	12.5	10.1	31.9	
Hypertension (%)	23.2	20.6	43.7	<.001
Diabetes mellitus (%)	4.1	3.2	11.4	<.001
Firearm noise exposure ( <i>Exposed %</i> )	7.5	6.8	13.3	0.001
Recreation noise exposure ( <i>Exposed %</i> )	26.0	25.6	29.3	0.203

<sup>a</sup>Hearing loss was defined as pure tone average at speech frequencies > 25 dB.

<sup>b</sup>Survey *t*-test (age-adjusted) for continuous variables and survey (Rao-Scott) Chi-square test for categorical variables were used.

<sup>c</sup>Pure tone average at speech frequencies at 0.5, 1, 2, and 4 kHz.

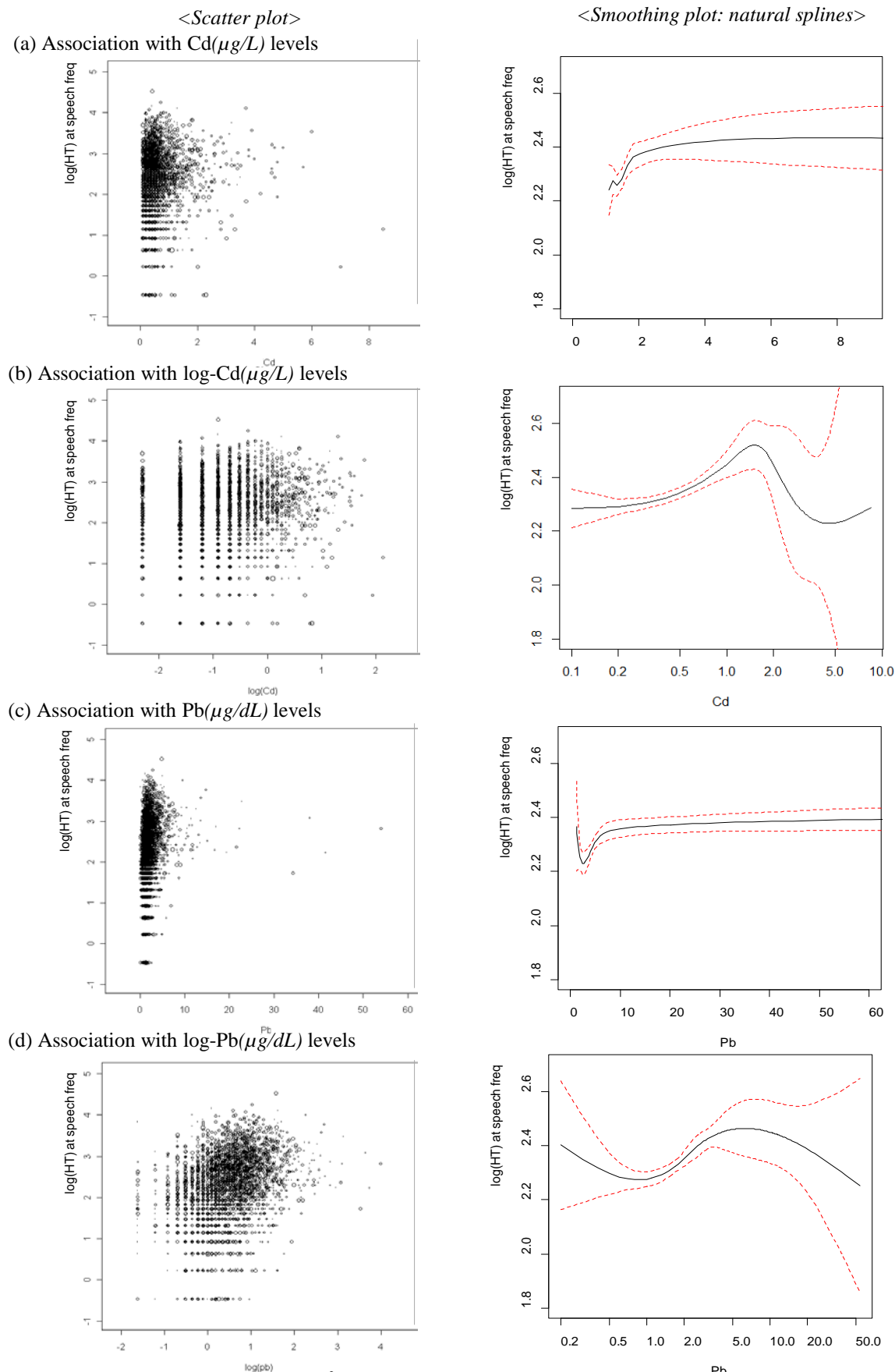
<sup>d</sup>Occupation noise (1 < O\*NET noise scale < 5).

**Supplemental Material, Table S3.** Percent change (95% CIs) of hearing thresholds (dB) by contribution of different variables in a multiple linear regression

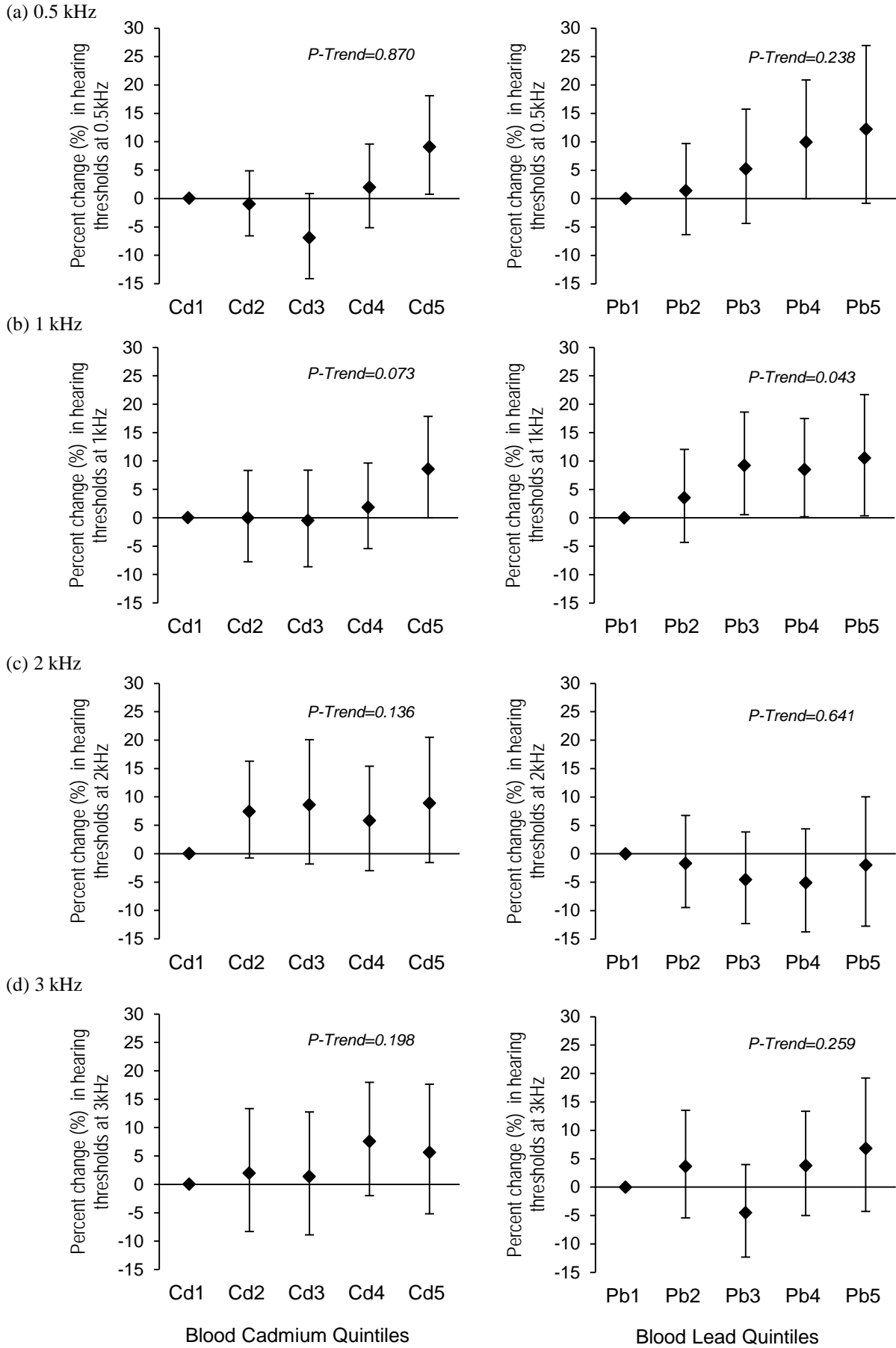
Variables	No.	Regression model <sup>a</sup>
Cadmium Quintile ( <i>ug/L</i> )		
Quintile 1 (0.10-0.20)	1013	0 (Reference)
Quintile 2 (0.30-0.30)	553	-1.22 (-8.86, 7.07)
Quintile 3 (0.40-0.40)	581	1.68 (-5.60, 9.53)
Quintile 4 (0.50-0.70)	785	6.69 (-1.48, 15.53)
Quintile 5 (0.80-8.50)	690	13.78 (4.55, 23.82)
Lead Quintile ( <i>ug/dL</i> )		
Quintile 1 (0.20-0.80)	629	0 (Reference)
Quintile 2 (0.90-1.30)	842	-0.5 (-9.94, 9.93)
Quintile 3 (1.40-1.80)	679	6.51 (-3.76, 17.89)
Quintile 4 (1.90-2.70)	734	10.22 (-0.40, 21.97)
Quintile 5 (2.80-54.00)	738	18.63 (7.35, 31.09)
O*NET Noise at longest job ( <i>unit score change</i> )		7.15 (1.48, 13.13)
Age ( <i>unit year change</i> )		1.38 (0.05, 2.73)
Age <sup>2</sup> ( <i>unit year change</i> )		0.01 (0.00, 0.03)
Body mass index ( <i>10 wtkg/htm change</i> )		1.23 (0.85, 1.61)
Sex		
Male	1729	0 (Reference)
Female	1969	-18.35 (-22.54, -13.94)
Race ethnics		
Non-Hispanic White	1827	0 (Reference)
Non-Hispanic Black	750	-11.88 (-17.96, -5.36)
Mexican American	805	-6.20 (-14.25, 2.60)
Other	316	0.78 (-8.49, 11.00)
Education		
< High School	974	0 (Reference)
High School	849	-9.78 (-16.99, -1.95)
> High School	1875	-19.81 (-24.20, -15.16)
Ototoxic medication		
No	3132	0 (Reference)
Yes	566	2.53 (-4.88, 10.52)
Cumulative cigarette packyears		
Never	2105	0 (Reference)
<20	1183	-1.80 (-8.96, 5.92)
≥20	410	4.46 (-4.45, 14.30)
Current dx of hypertension		
No	2713	0 (Reference)
Yes	985	-1.01 (-5.92, 4.16)
Current dx of diabetes mellitus		
No	3485	0 (Reference)
Yes	213	19.87 (8.93, 31.91)
Noise Exposure at firearm		
No	3468	0 (Reference)
Yes	230	10.19 (-0.52, 22.05)
Noise Exposure at recreation		
No	2844	0 (Reference)
Yes	854	3.43 (-4.01, 11.46)

<sup>a</sup>Regression model was adjusted for age, age<sup>2</sup>, bmi, sex, race/ethnicity, ototoxic medication, cumulative cigarette packyears, current dx of hypertension, current dx of diabetes, and occupation, recreation, and firearm noise. Cadmium models were further adjusted for lead; lead models were further adjusted for cadmium.

**Supplemental Material, Figure S1.** Graphical distribution of blood cadmium and lead with log-transformed hearing thresholds at speech frequencies (the average of thresholds at 0.5, 1, 2, and 4 kHz)

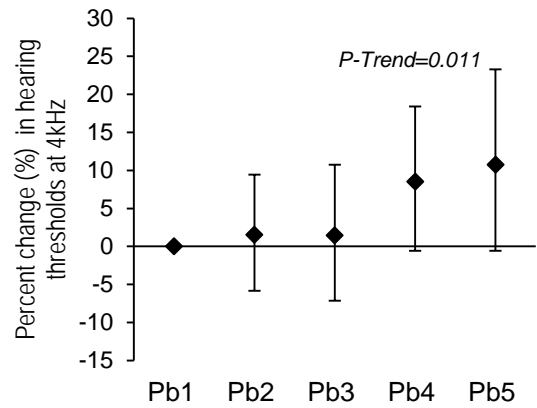
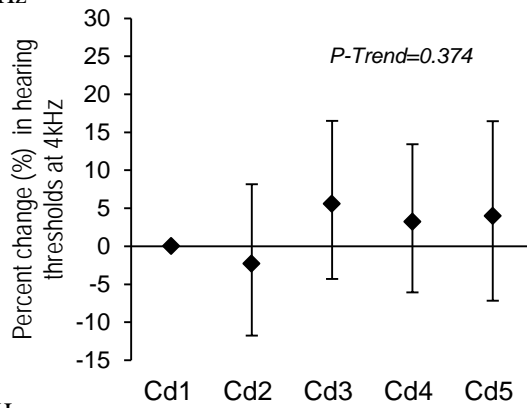


**Supplemental Material, Figure S2.** Percent change (%) of hearing thresholds (dB) by blood cadmium and lead quintiles at each frequency from 0.5 kHz to 8 kHz

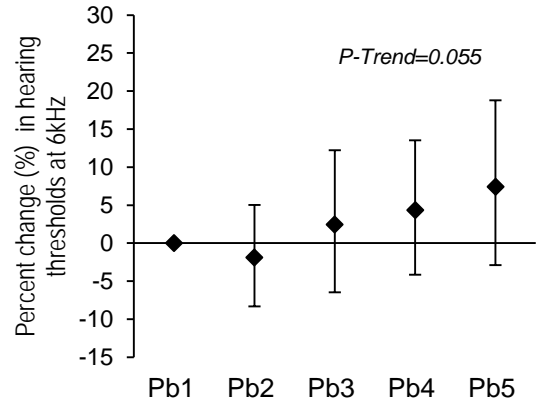
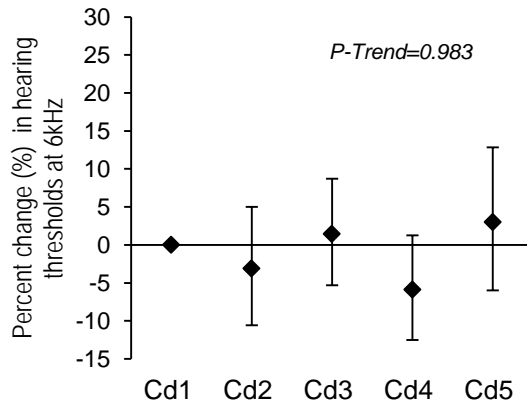


**Supplemental Material, Figure S2 (cont.).** Percent change (%) of hearing thresholds (dB) by blood cadmium and lead quintiles at each frequency from 0.5 kHz to 8 kHz

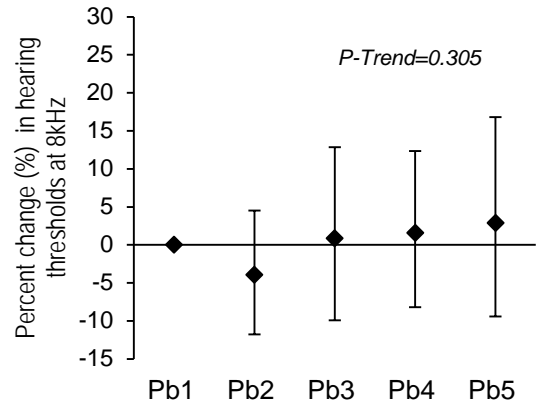
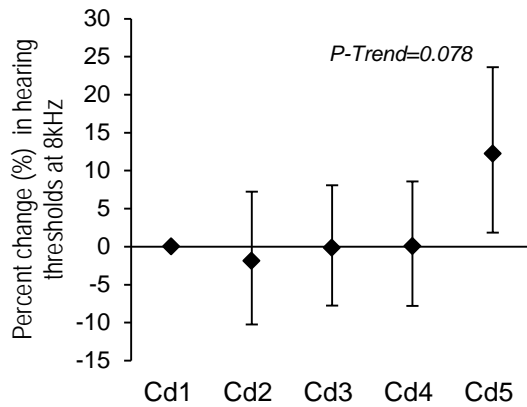
(e) 4 kHz



(f) 6 kHz



(g) 8 kHz



Blood Cadmium Quintiles

Blood Lead Quintiles

Regression models were adjusted for age, age<sup>2</sup>, body mass index, sex, race/ethnicity, education, ototoxic medication, cumulative cigarette pack-years, hypertension, diabetes, occupation noise, recreation noise, firearm noise. Cadmium models were further adjusted for lead; lead models were further adjusted for cadmium.